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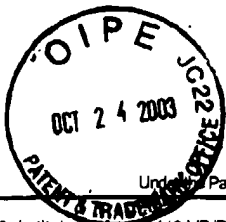
Substitute for form 1449A/B/PTO				Complete if Known	
				Application Number	10/615,716
INFORMATION DISCLOSURE STATEMENT BY APPLICANT (Use as many sheets as necessary)				Filing Date	July 8, 2003
				First Named Inventor	Mladen MERCEP
				Art Unit	TBA
				Examiner Name	TBA
				Attorney Docket Number	03818/100L650-US1
Sheet	1	of	4		

U.S. PATENT DOCUMENTS					
Examiner Initials*	Cite No. ¹	Document Number	Publication Date MM-DD-YYYY	Name of Patentee or Applicant of Cited Document	Pages, Columns, Lines, Where Relevant Passages or Relevant Figures Appear
		Number-Kind Code ² (if known)			
4	1.	6,297,260	10/02/2001	Bandarage et al.	
4	2.	4,710,495	12/01/1987	Bodor	
4	3.	6,402,733	06/11/2002	Daugherty	
4	4.	6,273,086	08/14/2001	Ohki et al.	
4	5.	6,228,346	05/08/2001	Zhang et al.	
4	6.	5,747,467	05/05/1988	Agouridas et al.	
4	7.	4,474,768	10/02/1984	Bright	

FOREIGN PATENT DOCUMENTS						
Examiner Initials*	Cite No. ¹	Foreign Patent Document	Publication Date MM-DD-YYYY	Name of Patentee or Applicant of Cited Document	Pages, Columns, Lines, Where Relevant Passages or Relevant Figures Appear	T ³
		Country Code ³ -Number ⁴ -Kind Code ⁵ (if known)				
4	8.	WO 94/13690	06/23/1992	Rhone-Poulenc Rorer Limited		
4	9	WO 94/14834	07/07/1994	Rhone-Poulenc Rorer Limited		
4	10.	WO 92/13873	08/20/1992	Aktiebolaget Astra		
4	11.	WO 92/13872	08/20/1992	Aktiebolaget Astra		
4	12.	WO 00/42055	07/20/2000	Zambon Group S.P.A.		
4	13.	EP 0283055	08/29/1990	Sour Pliva farmaceutska		
4	14.	EP 0775489	05/28/1997	Taisho Pharmaceutical Co. Ltd.		
4	15.	EP 0771564	05/07/1997	Taisho Pharmaceutical Co. Ltd.		
4	16.	WO 97/41255	11/06/1997	Massachusetts Institute of Technology		
4	17.	EP 00680967	10/14/1998	Hoechst Marion Roussel		
4	18.	WO 99/51616	10/14/1999	Pliva, Farmaceutska		
4	19.	EP 0984019	03/08/2000	Pfizer Products Inc.		
4	20.	WO 98/56801	12/17/1998	Pfizer Products Inc.		
4	21.	WO 94/14834	07/07/1994	Rhone-Poulenc Rorer Limited		
4	22.	EP 0984019 A1	03/08/2000	Pfizer Products Inc.		
4	23.	WO 98/56801	12/17/1998	Pfizer Products Inc.		

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Examiner Signature	<i>4. Patel</i>	Date Considered	<i>8/15/03</i>
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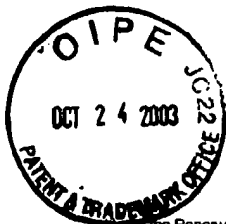
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NON PATENT LITERATURE DOCUMENTS			
Examiner Initials	Cite No. ¹	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published.	T ²
Gy	24.	Gladue R. P. et al., "In Vitro and In Vivo Uptake of Azithromycin (CP-62,993) by Phagocytic Cells: Possible Mechanism of Delivery and Release at Sites of Infection," <i>Antimicrob. Agents and Chemother.</i> , 33, 1989, 277-282	
Gy	25.	Olsen K. M. et al., "Intrapulmonary Pharmacokinetics of Zithromycin in Healthy Volunteers Given Five Oral Doses," <i>Antimicrob. Agents and Chemother.</i> , 40, 1996, 2582-2585	
Gy	26.	Mikasa, K. et al., "The anti-inflammatory effect of erythromycin in zymosan-induced peritonitis of mice," <i>J. Antimicrob. Chemother.</i> , 30, 1992, 339-348	
Gy	27.	"Discussion, Genomic organization of axolotl 1g genes," <i>J. Immunol.</i> , 159, 1997, 3395-4005	
Gy	28.	Takizawa, H. et al., "Erythromycin Modulates IL-8 Expression in Normal and Inflamed Human Bronchial Epithelial Cells," <i>Am. J. Respir. Crit. Care Med.</i> , 156, 1997, 266-271	
Gy	29.	Labro, M.T., "Anti-inflammatory activity of macrolides: a new therapeutic potential?" <i>J. Antimicrob. Chemother.</i> , 41, 1998, 37-46	
Gy	30.	Cazzola, M., et al., "Potential role of macrolides in the treatment of asthma," <i>Mondaldi Arch. Chest Dis.</i> , 55, 2000, 231-236	
Gy	31.	Avila, P.C. et al., "Macrolides, asthma, inflammation, and infection," <i>Ann. Allergy Asthma Immunol.</i> , 84, 2000, 565-568	
Gy	32.	Amayasu, H. et al., "Clarithromycin suppresses bronchial hyperresponsiveness associated with eosinophilic inflammation in patients with asthma," <i>Ann. Allergy, Asthma & Immunol.</i> , 84, 2000, 594-598	
Gy	33.	Shoji, T. et al., "Anti-inflammatory effect of roxithromycin in patients with aspirin-intolerant asthma," <i>Clin. Exp. Allergy</i> , 29, 1999, 950-956	
Gy	34.	Griffith, E.C., et al., "Yeast Three-Hybrid System for Detecting Ligand-Receptor Interactions," <i>Methods in Enzymology</i> , 328m 2000, 89-110	
Gy	35.	Denis A. et al., "Synthesis and Antibacterial Activity of HMR 36K47, A New Ketolide Highly Potent Against Erythromycin-Resistant and Susceptible Pathogens," <i>Bioorg. & Med. Chem. Lett.</i> , 9, 1999, 3075-3080	
Gy	36.	Agouridas C. et al., "Synthesis and Antibacterial Activity of Ketolides (6-O-Methyl-3-oxoerythromycin Derivatives): A New Class of Antibacterials Highly Potent against Macrolide-Resistant and -Susceptible Respiratory Pathogens," <i>J. Med. Chem.</i> , 41, 1998, 4080-4100	
Gy	37.	Sun, Or Y. et al. <i>J. Med. Chem.</i> 2000, 43, 1045-1049	
Gy	38.	McFarland, J. W. et al., "Repromicin Derivatives with Potent Antibacterial Activity against <i>Pasteurella multocida</i> ," <i>J. Med. Chem.</i> , 50, 1997, 1041-1045	
Gy	39.	Denis A. et al., "Synthesis of 6-O-Methyl-Azithromycin and Its Ketolide Analogue via Beckmann Rearrangement of 9(E)-6-O-Methyl-Erythromycin Oxime," <i>Bioorg. & Med. Chem. Lett.</i> , 8, 1998, 2427-2432	
Gy	40.	Lartey et al., "Synthesis of 4"-Deoxy Motilides: Identification of a Potent and Orally Active Prokinetic Drug Candidate, <i>J. Med. Chem.</i> , 38, 1998, 1793-1798	

Examiner Signature	<i>G. P. Kelly</i>	Date Considered	8/5/04
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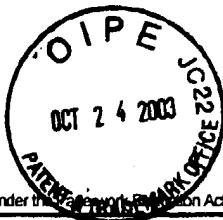
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				Art Unit	TBA
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Sheet	3	of	4	Attorney Docket Number	03818/100L650-US1

41.	Kirst, H.A. et al., "34. Metabolism of macrolides," Bryskier, A. J. et al., Ed. <i>Macrolides, Chemistry, Pharmacology and Clinical Use</i> ; Bryskier, Arnette Blackwell: Paris, 1993; pp 485-491
42.	Ma, Z. et al., "Discovery and Development of Ketolides as a New Generation of Macrolide Antimicrobial Agents," <i>Current Medicinal Chemistry - Anti-Infective Agents</i> , 1, 2002, 15-34
43.	Pascual A. et al., "Uptake and intracellular activity of ketolide HMR 3647 in human phagocytic and non-phagocytic cells," <i>Clin. Microbiol. Infect.</i> , 7, 2001, 65-69
44.	Hand, W. L. et al., "Characteristics and mechanisms of azithromycin accumulation and efflux in human polymorphonuclear leukocytes," <i>Int. J. Antimicrob. Agents</i> , 18, 2001, 419-425
45.	Amsden, G. W., "Advanced-generation macrolides: tissue-directed antibiotics," <i>Int. J. Antimicrob. Agents</i> , 18, 2001, 11-15
46.	Johnson, J. D. et al., "Antibiotic uptake by alveolar macrophages," <i>J. Lab. Clin. Med.</i> , 95, 1980, 429-439
47.	Wildfeuer, A. et al., "Uptake of Azithromycin by Various Cells and Its Intracellular Activity under In Vivo Conditions," <i>Antimicrob. Agents Chemother.</i> , 40, 1996, 75-79
48.	Scorneaux, B. et al., "Intracellular Accumulation, Subcellular Distribution, and Efflux of Tilimicosin in Chicken Phagocytes," <i>Poult. Sci.</i> , 77, 1998, 1510-1521
49.	Mtairag, E. M. et al., "Investigation of dirithromycin and erythromyclamine uptake by human neutrophils in vitro," <i>J. Antimicrob. Chemother.</i> 33, 1994, 523-536
50.	Anderson R. et al., "An in-vitro evaluation of the cellular uptake and intraphagocytic bioactivity of clarithromycin (A-56268, TE-031), a new macrolide antimicrobial agent," <i>J. Antimicrob. Chemother.</i> , 22, 1988, 923-933
51.	Tasaka, Y. et al., "Rokitamycin Uptake by Alveolar Macrophages," <i>Jpn. J. Antibiot.</i> 41, 1988, 836-840
52.	Harf, R. et al., "Spiramycin uptake by alveolar macrophages," <i>J. Antimicrob. Chemother.</i> , 22, 1988, 135-140
53.	Suzuki, T. et al., "General and facile method for determination of configuration of steroid-17-yl-methyl glycolates at C-20 based on kinetic examination," <i>Chem. Soc., Perkin Trans. 1</i> , 1998, 3831-3836
54.	McLean, H.M. et al., "Novel Fluorinated Antiinflammatory Steroid with Reduced Side Effects: Methyl 9 α -Fluoroprednisolone-16-carboxylate," <i>J. Pharm. Sci.</i> 1994, 83, 476-480
55.	Little, R.J. et al., "Soft Drugs Based on Hydrocortisone: The Inactive Metabolite Approach and Its Application to Steroidal Antiinflammatory Agents," <i>Pharm. Res.</i> , 16, 1999, 961-967
56.	Kertesz, D.J. et al., "Thiol Esters from Steroid 17 β -Carboxylic Acids: Carboxylate Activation and Internal Participation by 17 α -Acylates," <i>J. Org. Chem.</i> , 51, 1986, 2315-2328
57.	Phillipps, G. et al., "Synthesis and Structure - Activity Relationships in a Series of Antiinflammatory Corticosteroid Analogues, Halomethyl Androstane-17 β -carbothioates and 17 β -carbosenoates," <i>J. Med. Chem.</i> 37, 1994, 3717-3729
58.	Bright, G.M. et al., "Synthesis, In Vitro and In Vivo Activity of Novel 9-Deoxy-9a-AZA-9a-Homoerythromycin A Derivatives; A new Class of Macrolide Antibiotics, the Azalides" <i>J. Antibiot.</i> , 41, 1998, 1029-1047
59.	Costa, A.M. et al., "Hybrids of macrolides and nucleobases or nucleosides," <i>Tetrahedron Letters</i> , 41, 2000, 3371-3375

Examiner Signature	<i>G. R. K.</i>	Date Considered	<i>3/1/04</i>
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60.	Newman, S.P. et al., "Evaluation of jet nebulisers for use with gentamicin solution," <i>Thorax</i> , 40, 1985, 671-676
61.	Berenberg, M.J. et al., "Comparison of Metered-Dose Inhaler Attached to an Aerochamber with an Updraft Nebulizer for the Administration of Metaproterenol in Hospitalized Patients," <i>J. Asthma USA</i> , 22, 1985, 87-92

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